

Listing of Claims

1-29. (Canceled)

30. (Currently Amended) A method for identifying a molecule that binds a known target receptor in a cell, comprising:

- (a) forming a screening molecule by covalently bonding the molecule to be identified to a moiety capable of selectively binding to and selectively forming a covalent bond with a penicillin-binding-protein ("PBP") or a thymidine synthase ("TS") enzyme receptor domain;
- (b) introducing the screening molecule into a cell culture comprising cells that express a first fusion protein of a LexA DNA-binding domain fused to [[a]] the known target receptor domain, a second fusion protein which comprises (i) [[a]] the penicillin-binding-protein ("PBP") or [[a]] the thymidine synthase ("TS") enzyme receptor domain ~~capable of binding to and forming the~~ ~~covalent bond with the screening molecule~~ and (ii) a B42 transcription activation domain, and a reporter gene wherein expression of the reporter gene is conditioned on the proximity of the first fusion protein to the second fusion protein;
- (c) permitting the screening molecule to bind to the first fusion protein and to the second fusion protein, bringing the two fusion proteins in to proximity so as to activate the expression of the reporter gene,
- (d) selecting the cell that expresses the reporter gene; and
- (e) identifying the molecule that binds the known target receptor wherein cellular expression of the reporter gene indicates that the molecule is able to bind to the known target receptor.

31. (Previously Presented) The method of claim 30, wherein the cell is selected from the group consisting of insect cells,

*Enkro*  
*11/5/08*  
*JL*

yeast cells, mammalian cells, and their lysates.

32. (Canceled).

33. (Canceled).

34. (Canceled).

35. (Previously Presented) The method of claim 30, wherein the PBP is the *Streptomyces R61 PBP*.

36. (Currently Amended) The method of claim 30, wherein the molecule to be identified is obtained from a combinatorial library.

37. (Previously Presented) The method of claim 30, wherein the steps (b)-(c) of the method are iteratively repeated in the presence of a preparation of random molecules for competitive binding with the screening molecule so as to identify a molecule capable of competitively binding the known target receptor.

38. (Currently Amended) A method for identifying a target receptor as being able to bind to a ligand, comprising:

- (a) providing a screening molecule having a ligand covalently bonded to [(a)] a moiety capable of selectively binding to and selectively forming a covalent bond with a penicillin-binding-protein ("PBP") or a thymidine synthase ("TS") enzyme second receptor;
- (b) introducing the screening molecule into a cell which expresses  
a first fusion protein of a LexA DNA-binding domain fused to the target receptor,

a second fusion protein which comprises

(i) a penicillin-binding-protein ("PBP") or a thymidine synthase ("TS") enzyme second receptor domain capable of binding to and forming the covalent bond with the screening molecule and (ii) and a B42 transcription activation domain, and

a reporter gene wherein expression of the reporter gene is conditioned on the proximity of the first fusion protein to the second fusion protein;

(c) permitting the screening molecule to bind to the first fusion protein and to the second fusion protein so as to activate the expression of the reporter gene,

(d) selecting which cell expresses the unknown target receptor; and

(e) identifying the unknown target receptor wherein cellular expression of the reporter gene indicates that the target receptor is able to bind to the ligand.

39. (Previously Presented) The method of claim 38, wherein the target receptor is encoded by a DNA from the group consisting of genomic DNA, cDNA and synthetic DNA.

40. (Previously Presented) The method of claim 38, wherein the ligand has a known biological function.

41-56. (Canceled)

57. (Previously Presented) The method of claim 30, wherein the chemical inducer of dimerization comprises a cephem or a fluorouracil moiety.

58. (Previously Presented) The method of claim 38, wherein the

Applicant: Virginia W. Cornish

Serial No.: 10/056,874

Filed: January 24, 2002

Page 5

substrate comprises a cephem or a fluorouracil moiety.